Internati pplication No PCT/GB 02/04633

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N15/11 C07K14/47 C12N15/67 A61K38/16 A61K47/48 C12N5/10 A01K67/027 C12N15/82

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12N C07K A61K A01K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, EMBASE, MEDLINE

43
5, -16, ,38
2, 10, -35,
, 39–43
5, -16, ,38

Further documents are fisted in the continuation of box C.	Y Palent family members are listed in annex.
Special categories of cited documents: 'A' document delining the general state of the art which is not considered to be of particular relevance 'E' earlier document but published on or after the international filling date 'L' document which may throw doubts on priority ctaim(s) or which is cited to establish the publication date of another clastion or other special reason (as specified) 'O' document referring to an oral disclosure, use, exhibition or other means 'P' document published prior to the international filing date but later than the priority date calmed	 "T" later document published after the International filling date or priority date and not in conflict with the application but died to understand the principle or theory underlying the invention. "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone. "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents; such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Dale of the adual completion of the International search 12 February 2003	Date of mailing of the international search report 26/02/2003
Name and mailing address of the ISA European Palant Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijawijk Tol. (+31-70) 340-2040, Tx. 31 651 epo ni, Fax. (+31-70) 340-3018	Authorized onloar Griesinger, I

Internatio pplication No PCT/GB 02/04633

Y WO 01 59094 A (SA STUDI) 16 August	on, where appropriate, of the relevant passages	Relevant to claim No.
Y WO 01 59094 A (SA STUDI) 16 August		Relevant to claim No.
STUDI) 16 August	ALK THST FOR BIOLOGICAL	
abstract	2001 (2001-08-16)	1-43
antisense oligon EUROPEAN JOURNAL BIOPHARMACEUTICS PUBLISHERS B.V., vol. 50, no. 1, 3 pages 101-119, XF ISSN: 0939-6411 page 106, right-h	OF PHARMACEUTICS AND , ELSEVIER SCIENCE AMSTERDAM, NL, 3 July 2000 (2000-07-03),	1-43
transcriptional r GENES AND DEVELOP LABORATORY PRESS,	one acetylation and regulatory mechanisms" PMENT, COLD SPRING HARBOR NEW YORK, US, ages 599-606, XP002157338	

nal application No. PCT/GB 02/04633

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inter	mational Search Report has not been established in respect of certain dalms under Article 17(2)(a) for the following reasons:
	Claims Nos.: 37 (partially) because they relate to subject matter not required to be searched by this Authority, namely: see FURTHER INFORMATION sheet PCT/ISA/210
	. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful international Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
	Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This inter	mational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable daims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this international Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4	No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the dalms; it is covered by claims Nos.:
Р едзік	on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

International Application No. PCT/GB 02 D4633

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 1-23, 27, and 42 could be interpreted as being directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound.

Continuation of Box 1.1

Claims Nos.: 37 (partially)

The term "animal" could be interpreted as including humans. However, transgenic humans are not considered patentable (Rule 9.1 PCT).

ERIC PUTTER CLARKSON

INTERNATIONAL SEARCH REPORT

nf tion on patent family members

Internatio pplication No PCT/GB 02/04633

Patent document died in search report		Publication date		Patent family member(6)	Publication date
WO 0102019	A	11-01-2001	AU	5556600 A	22-01-2001
			EP	1190073 A2	27-03-2002
			WO	0102019 A2	11-01-2001
			GB	2367555 A	10-04-2002
WO 9911809	A	11-03-1999	AU	737829 B2	30-08-2001
MO 3311003	^	00 ,	AU	8877698 A	22-03-1999
			BR	9811744 A	02-01-2002
			CA	2301157 A1	11-03-1999
			EP	1009847 A1	21-06-2000
			WO	9911809 A1	11-03-1999
		•	ĴΡ	2001514858 T	18-09-2001
			NZ	503007 A	24-11-2000
WO 0159094	A	16-08-2001	AU	3696101 A	20-08-2001
MO 0103034	^	10 00 2001	WO	0159094 A2	16-08-2001
			ÜS	2002022021 A1	21-02-2002

PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY	PCT					
To: ERIC POTTER CLARKSON Attn. Miles, John S. Park View House 58 The Ropewalk Nottingham NG1 5DD UNITED KINGDOM	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION (PCT Rule 44.1)					
81	Date of mailing (day/month/year) 26/02/2003					
Applicants or agents file reference GENY/P27203PC	FOR FURTHER ACTION See paragraphs 1 and 4 below					
International application No. PCT/GB 02/04633	International filing date (day/month/year) 11/10/2002					
Applicant						
GENE EXPRESSION TECHNOLOGIES LIMITED						
The applicant is hereby notified that the International Search Report has been established and is transmitted herewith. Fiting of amendments and statement under Article 19: The applicant is entitled, If he so wishes, to amend the claims of the International Application (see Rule 46): When? The time limit for filling such amendments is normally 2 months from the date of transmittal of the International Search Report; however, for more details, see the notes on the accompanying sheet.						
Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41-22) 740.14.35						
For more detailed instructions, see the notes on the according	ompanying sheet					
2. The applicant is hereby notified that no International Searce Article 17(2)(a) to that effect is transmitted herewith.	h Report will be established and that the declaration under					
With record to the project against payment of (an) additi	onal fee(s) under Rule 40.2, the applicant is notified that:					
3. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the lexts of both the protest and the decision thereon to the designated Offices.						
no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made.						
4. Further action(s): The applicant is reminded of the following:	III ha autiliahad hu sha labagashapal Duragu					
Shortly after 16 months from the priority date, the International a if the applicant wishes to avoid or postpone publication, a notic priority claim, must reach the International Bureau as provided completion of the technical preparations for international public	t in Rules 90 bls.1 and 90 bls.3, respectively, before the ration.					
Within 19 months from the priority date, a demand for internation wishes to postpone the entry into the national phase until 30 m	ionins nom the phoney date in some or have a					
Within 20 months from the priority date, the applicant must perform before all designated Offices which have not been elected in the priority date or could not be elected because they are not bour	THE CENTIAND OF MY & MAICH CHESCOTT WISHING TO THE CONTROL OF THE					
Name and mailing address of the International Searching Authority	Authorized officer					

Catriona Cleere

European Patent Office, P.B. 5818 Patentiaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international polication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the daims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Flute 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been fis filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended, it must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- (Where originally there were 48 claims and after amendment of some claims there are 51):
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
 claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- 3. [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended, it must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international proliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

	Applicant's or agent's file reference GENY/P27203PC	FOR FURTHER see Notification of (Form PCT/ISA/2	f Transmittal of International Search Report 20) as well as. where applicable, item 5 below.
	International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
	PCT/GB 02/04633	11/10/2002	11/10/2001
	Applicant	-	
	GENE EXPRESSION TECHNOLOG	IES LIMITED	
	This International Search Report has been according to Article 18. A copy is being tra	prepared by this International Searching Authorsmitted to the International Bureau.	nority and is transmitted to the applicant
	This International Search Report consists It is also accompanied by	of a total of6 sheets, a copy of each prior art document cited in this	report
Ī	1. Basis of the report		
	 a. With regard to the language, the language in which it was filed, unle 	nternational search was carried out on the basess otherwise indicated under this item.	sls of the International application in the
	Authority (Rule 23.1(b)).	as carried out on the basis of a translation of t	
	 With regard to any nucleotide and was carried out on the basis of the 	d <mark>/or amino acid sequence disclosed in the in</mark> e sequence listing :	temational application, the international search
-		nal application in written form.	
	liled together with the inter	mational application in computer readable form	n.
	furnished subsequently to	this Authority in written form.	
- [this Authority in computer readble form.	
	international application as		
	the statement that the info furnished	rmation recorded in computer readable form is	identical to the written sequence listing has been
1	2. X Certain claims were four	nd unsearchable (See Box I).	
	3. Unity of invention is lack	ting (see Box II).	
	4. With regard to the title,		
-	the text is approved as sub	omitted by the applicant.	
ł	the text has been establish	ned by this Authority to read as follows:	OF TOOMING BOTTOE AND A DECIDATO
	CONTROL OF GENE EXPRES	SION USING A COMPLEX OF AN	OLIGONUCLEOTIDE AND A REGULATO
	5. With regard to the abstract.		
1	the text is approved as sub		
	the text has been establish within one month from the	ned, according to Rule 38.2(b), by this Authorit date of mailing of this international search rep	y as it appears in Box III. The applicant may, on, submit comments to this Authority.
	6. The figure of the drawings to be publis		1
	as suggested by the applic		None of the figures.
	because the applicant faile		
	because this figure better o	characterizes the invention.	

International application No.

INTERNATIONAL SEARCH REPORT

PCT/GB 02/04633

Box III TEXT OF THE ABSTRACT (Continuation of item 5 of the first sheet)

A method for suppressing the expression of a selected gene in a cell the method comprising introducing into the cell a molecule comprising (1) a nucleic acid binding portion which binds to a site or associated with the selected gene which site is present in a genome and (2) an expression repressor portion, wherein the nucleic acid binding portion comprises an oligonucleotide or oligonucleotide mimic or analogue, and wherein the repressor portion comprises a polypeptide or peptidomimetic. Molecules for use in the methods of the invention are provided. The repressor may be a portion of a histone deacetylase or DNA methylase or polypeptide capable of recruiting a histone deacetylase or DNA methylase.

International Application No PCT/GB 02/04633

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N15/11 C07K14/47

C12N5/10

A01K67/027

C12N15/67 C12N15/82

A61K38/16

A61K47/48

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12N C07K A61K A01K

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Documentation scarched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, EMBASE, MEDLINE

Category •	Citation of document, with Indication, where appropriate, of the relevant passages	Relevant to claim No.
Х	WO 01 02019 A (IMP COLLEGE INNOVATIONS LTD; ALI SIMAK (GB); BULUWELA LAKJAYA (GB))	1-43
Υ	11 January 2001 (2001-01-11) abstract; claims 1-8,16-24,45-49	3-5, 11-16, 36,38
X	WO 99 11809 A (IMP COLLEGE INNOVATIONS LTD;CRISANTI ANDREA (GB)) 11 March 1999 (1999-03-11)	1,2, 6-10, 17-35, 37,39-43
	page 10, line 1-13 page 12, line 11 -page 13, line 3 page 16, line 1-14	
Y	page 10, Time 1 14	3-5, 11-16, 36,38
	 -/	

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

- Special categories of cited documents:
- "A" document defining the general state of the art which is not considered to be of particular relevance
- *E* earlier document but published on or after the international filling date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- document referring to an oral disclosure, use, exhibition or other means
- document published prior to the International Tiling date but later than the priority date claimed
- The later document published after the International Illing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the Invention
- 'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- '&' document member of the same patent family

Date of mailing of the international search report

Date of the actual completion of the international search

12 February 2003

Name and mailing address of the ISA European Palent Office, P.B. 5818 Falentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl. Fax: (+31-70) 340-3016

26/02/2003

Authorized officer

Griesinger, I

Form PCT/ISA/210 (second shoot) (July 1992)

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International Application No
PCT/GB 02/04633

		PCT/GB 02	/ U4033
	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		Relevant to claim No.
Category *	Citation of document, with indication, where appropriate, of the relevant passages		
1	WO 01 59094 A (SALK INST FOR BIOLOGICAL STUDI) 16 August 2001 (2001-08-16) abstract		1-43
(LEBEDEVA I ET AL: "Cellular delivery of antisense oligonucleotides" EUROPEAN JOURNAL OF PHARMACEUTICS AND BIOPHARMACEUTICS, ELSEVIER SCIENCE PUBLISHERS B.V., AMSTERDAM, NL, vol. 50, no. 1, 3 July 2000 (2000-07-03), pages 101-119, XP004257182 ISSN: 0939-6411 page 106, right-hand column, paragraph 5 page 109, left-hand column, paragraph 2		1-43
1	STRUHL K: "histone acetylation and transcriptional regulatory mechanisms" GENES AND DEVELOPMENT, COLD SPRING HARBOR LABORATORY PRESS, NEW YORK, US, vol. 12, 1998, pages 599-606, XP002157338 ISSN: 0890-9369		
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Form PCT/ISA/210 (continuation of second sheet) (July 1992)

International Application No. PCT/GB 02 \(D4633 \)

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box 1.1

Although claims 1-23, 27, and 42 could be interpreted as being directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound.

Continuation of Box I.1

Claims Nos.: 37 (partially)

The term "animal" could be interpreted as including humans. However, transgenic humans are not considered patentable (Rule 9.1 PCT).

International application No. PCT/GB 02/04633

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
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Remark	on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

Information on patent family members

International Application No PCT/GB 02/04633

Patent document dted in search report		Publication date		Patent family member(s)	Publication date
WO 0102019	A	11-01-2001	AU EP	5556600 A 1190073 A2	22-01-2001 27-03-2002
			MO	0102019 A2	11-01-2001
			GB	2367555 A	10-04-2002
WO 9911809	A	11-03-1999	AU	737829 B2	30-08-2001
NO JJIIOOJ	, ,		ΑU	8877698 A	22-03-1999
			BR	9811744 A	02-01-2002
			CA	2301157 A1	11-03-1999
			EP	1009847 A1	21-06-2000
			WO	9911809 A1	11-03-1999
			JP	2001514858 T	18-09-2001
			NZ	503007 A	24-11-2000
WO 0159094	Α	16-08-2001	AU	3696101 A	20-08-2001
WO 0107074			WO	0159094 A2	16-08-2001
			US	2002022021 Al	21-02-2002

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Internati pplication No PCT/GB 02/04633

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N15/11 C07K14/47 C12N5/10 A01K67/027 A61K38/16 C12N15/67 A61K47/48 A01K67/027 C12N15/82

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Mhimum documentation searched (classification system followed by classification symbols) IPC 7 C12N C07K A61K A01K

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Category °	Citation of document, with indication, where appropriate, of the	ne relevant passages	Relevant to claim No.
Х	WO 01 02019 A (IMP COLLEGE INN ;ALI SIMAK (GB); BULUWELA LAKJ 11 January 2001 (2001-01-11)		1-43
Y	abstract; claims 1-8,16-24,45-	49	3-5, 11-16, 36,38
X	WO 99 11809 A (IMP COLLEGE INN ;CRISANTI ANDREA (GB)) 11 March 1999 (1999-03-11) page 10, line 1-13	OVATIONS LTD	1,2, 6-10, 17-35, 37,39-43
Y	page 12, line 11 -page 13, lin page 16, line 1-14	e 3	3-5, 11-16, 36,38
	er documents are listed in the continuation of box C.	-/ X Patent family members are listed	in annëx.
"A" docume consider of filling docume which I citation other n	nt which may throw doubts on priority claim(s) or s cited to establish the publication date of another or other special reason (as specified) nt referring to an oral disclosure, use, exhibition or	 'T' later document published after the inte or priority date and not in conflict with cited to understand the principle or the invention 'X' document of particular relevance; the cannot be considered novel or cannot involve an inventive step when the do 'Y' document of particular relevance; the cannot be considered to involve an involve an involve and in document is combined with one or mornants, such combination being obvious in the art. '&' document member of the same patent 	the application but cory underlying the laimed invention be considered to current is taken alone laimed invention rentive step when the re other such docusis to a person skilled
Date of the a	ctual completion of the International search	Date of malling of the international sea	rch report
12	Prebruary 2003	26/02/2003	
vame and m	ailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Griesinger, I	

Internatio pplication No
PCT/GB 02/04633

Y WO 01 59094 A (SAL STUDI) 16 August 2 abstract Y LEBEDEVA I ET AL: antisense oligonud EUROPEAN JOURNAL OBIOPHARMACEUTICS, PUBLISHERS B.V., A vol. 50, no. 1, 3 pages 101-119, XPO ISSN: 0939-6411	where appropriate, of the relevant passages LK INST FOR BIOLOGICAL 2001 (2001-08-16) "Cellular delivery of cleotides" DF PHARMACEUTICS AND ELSEVIER SCIENCE AMSTERDAM, NL, July 2000 (2000-07-03),	Relevant to	
Y WO 01 59094 A (SAL STUDI) 16 August 2 abstract Y LEBEDEVA I ET AL: antisense oligonuc EUROPEAN JOURNAL C BIOPHARMACEUTICS, PUBLISHERS B.V., A vol. 50, no. 1, 3 pages 101-119, XPO ISSN: 0939-6411	"Cellular delivery of cleotides" DF PHARMACEUTICS AND ELSEVIER SCIENCE MSTERDAM, NL, July 2000 (2000-07-03),	1-	43
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antisense oligonud EUROPEAN JOURNAL O BIOPHARMACEUTICS, PUBLISHERS B.V., A vol. 50, no. 1, 3 pages 101-119, XPO ISSN: 0939-6411	Cleotides" DF PHARMACEUTICS AND ELSEVIER SCIENCE AMSTERDAM, NL, July 2000 (2000-07-03),	1-	43
-page 109, left-ha	and column, paragraph 5 and column, paragraph 2		
GENES AND DEVELOPM LABORATORY PRESS,	egulatory mechanisms" MENT, COLD SPRING HARBOR		

Inter nal application No. PCT/GB 02/04633

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This international Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: 37 (partially) because they relate to subject matter not required to be searched by this Authority, namely: see FURTHER INFORMATION sheet PCT/ISA/210
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful international Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This international Searching Authority found multiple inventions in this international application, as follows:
1. As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 1-23, 27, and 42 could be interpreted as being directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound.

Continuation of Box I.1

Claims Nos.: 37 (partially)

The term "animal" could be interpreted as including humans. However, transgenic humans are not considered patentable (Rule $9.1\ PCT$).

Inf tion on patent family members

Internatio pplication No PCT/GB 02/04633

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 0102019	A	11-01-2001	AU	5556600 A	22-01-2001
			EP	1190073 A2	27-03-2002
			WO	0102019 A2	11-01-2001
			GB	2367555 A	10-04-2002
WO 9911809	A	11-03-1999	AU	737829 B2	30-08-2001
			AU	8877698 A	22-03-1999
			BR	9811744 A	02-01-2002
			CA	2301157 A1	11-03-1999
			EP	1009847 A1	21-06-2000
			WO	9911809 A1	11-03-1999
			JP	2001514858 T	18-09-2001
			NZ	503007 A	24-11-2000
WO 0159094	Α	16-08-2001	AU	3696101 A	20-08-2001
			WO	0159094 A2	16-08-2001
			US	2002022021 A1	21-02-2002

PATENT COOPERATION TREATY

PCT/GB02/04633

PCT

NOTICE INFORMING THE APPLICANT OF THE COMMUNICATION OF THE INTERNATIONAL APPLICATION TO THE DESIGNATED OFFICES

(PCT Rule 47.1(c), first sentence)

To:

MILES, John, S.
Eric Potter Clarkson
Park View House
58 The Ropewalk
Nottingham NG1 5DD
ROYAUME-UNI

IMPORTANT NOTICE

Date of mailing(day/month/year) 24 April 2003 (24.04.03)

Applicant's or agent's file reference GENY/P27203PC

International application No.

PCT/GB02/04633

International filing date(day/month/year) P.

Priority date(day/month/year)
11 October 2001 (11.10.01)

Applicant

GENE EXPRESSION TECHNOLOGIES LIMITED

11 October 2002 (11.10.02)

1. Notice is hereby given that the International Bureau has communicated, as provided in Article 20, the international application to the following designated Offices on the date indicated above as the date of mailing of this notice:

KP, KR, US

In accordance with Rule 47.1(c), third sentence, those Offices will accept the present notice as conclusive evidence that the communication of the international application has duly taken place on the date of mailing indicated above and no copy of the international application is required to be furnished by the applicant to the designated Office(s).

2. The following designated Offices have waived the requirement for such a communication at this time:

AE, AG, AL, AM, AP, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EA, EC, EE, EP, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OA, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW

The communication will be made to those Offices only upon their request. Furthermore, those Offices do not require the applicant to furnish a copy of the international application (Rule 49.1(a-bis)).

- Enclosed with this notice is a copy of the international application as published by the International Bureau on 24 April 2003 (24.04.03) under No. 03/033701
- 4. TIME LIMITS for filing a demand for international preliminary examination and for entry into the national phase

The applicable time limit for entering the national phase will, subject to what is said in the following paragraph, be 30 MONTHS from the priority date, not only in respect of any elected Office if a demand for international preliminary examination is filled before the expiration of 19 months from the priority date, but also in respect of any designated Office, in the absence of filing of such demand, where Article 22(1) as modified with effect from 1 April 2002 applies in respect of that designated Office. For further details, see PCT Gazette No. 44/2001 of 1 November 2001, pages 19926, 19932 and 19934, as well as the PCT Newsletter, October and November 2001 and February 2002 issues.

In practice, time limits other than the 30-month time limit will continue to apply, for various periods of time, in respect of certain designated or elected Offices. For regular updates on the applicable time limits (20, 21, 30 or 31 months, or other time limit), Office by Office, refer to the PCT Gazene, the PCT Newsletter and the PCT Applicant's Guide, Volume II, National Chapters, all available from WIPO's Internet site, at http://www.wipo.int/pct/enfindex.html.

For filing a demand for international preliminary examination, see the PCT Applicant's Guide, Volume VA, Chapter IX. Only an applicant who is a national or resident of a PCT Contracting State which is bound by Chapter II has the right to file a demand for international preliminary examination (at present, all PCT Contracting States are bound by Chapter II).

It is the applicant's sole responsibility to monitor all these time limits.

The applicant is hereby notified that, at the time of establishment of this Notice, the time limit under Rule 46.1 for making amendments under Article 19 has not yet expired and the International Bureau had received neither such amendments nor a declaration that the applicant does not wish to make amendments.

The International Bureau of WIPO 34, chemin des Colombeucs 1211 Geneva 20, Switzerland Authorized officer

Judith Zahra

Facsimile No.(41-22) 740.14.35

Telephone No.(41-22) 338.91.11



PCT

REQUEST

	For receiving Office use only	
International Applicati	on No.	
		-
International Filing Da	te	
Name of receiving Off	ice and "PCT International Appl	ication"

internation	al application	quests that the press be processed accor operation Treaty.	ent ding to		Applicant's	or agen	Office and "PCT International Application" I's file reference GENY/ P27203PC racters maximum)
	TTLE OF IN CONTROL	VENTION OF GENE EXP	RESSI	ON			
Box No. II A	PPLICANT			This person is	also inventor	7	
Name and address: (1) The address must incluse Box is the applicant's Six Gene Expression To 90 Fetter Lanc	de pustal code ute (that is, cou	and name of country ntry) of residence if no	. The cu	runtry of the add	ress indicated	nation. In thus	Telephone No.
London EC4A 1JP United Kingdom						į	Teleprinter No.
							Applicant's registration No. with the Office
State (that is, country,) of nationali	y: GB			State (that i.	s, count	ry) of residence: GB
This person is applicant for the purposes of:		all designated States	x	all designated St the United States			the United States the States indicated in the Supplemental Box
Box No. III F	URTHER A	PLICANT(S) AN	D/OR (FURTHER) II	NVENTOR(S	S)	
Name and address: (The address must inclust fine is the applicant's Note HART, Stephen Division of Medicit Imperial College So Room 601b, Cyclot Du Cane Road London W12 0NN United Kingdom	de possal code one (that is, cou one, Cancer (chool of Me	and name of country ntry) of residence if no Cell Biology Secti dicine, Hammerst	v. The co State of On	ountry of the ad residence is indic	dress indicated	gnation, I In this	This person is: applicant only applicant and inventor inventor only (If this check-box Is marked, do not fill in below) Applicant's registration No. with the Office
State (that is, country,	of nationalit	y: GB			State (that L	s, counti	ry) of residence: GB
This person is applicant for the purposes of:		all designated States		all designated St the United States		X	the United States the States indicated in the Supplemental Box
X Further applie	cants and/or (further) inventors a	re indic	ated on a contin	uation sheet.		
Box No. IV A	GENT OR C	OMMON REPRE	SENTA	ATIVE; OR A	DDRESS FO	R COR	RESPONDENCE
The person identified of the applicant(s) bet	below is here fore the comp	by/has been appoin etent International A	ted to a Luthorit	ct on behalf ics as:		X	agent common sepresentative
Name and address: (I The address must include Miles, John S	Family name f	ollowed by given nam			l official design	nation.	Telephone No. (0115) 9552211
Eric Potter Clarkson Park View House	n						Facsimile No. (0115) 9552201
58 The Ropewalk Nottingham NG1 5DD						ľ	Teleprinter No. 37540 Potter G
England							Agent's registration No. with the Office
		nce: Mark this chec					ative is/has been appointed and the pe sent.

Sheet No ..2.. FURTHER APPLICANT(S) AND/OR (FURTHER) INVENTOR(S) Continuation of Box No. III If none of the following sub-boxes is used, this sheet should not be included in the request. Name and address: (Family name followed by given name; for a legal entity, full official designation, This person is: The address must include postal code and name of country. The country of the address indicated in this Box is the opplicant's State (that is, country) of residence if no State of residence is indicated helow.) applicant only Division of Medicine, Cancer Cell Biology Section X applicant and inventor Imperial College School of Medicine, Hammersmith Campus inventor only (If this check-box is marked, Room 601b, Cyclotron Building do not fill in below) Du Cane Road Applicant's registration No. with the Office Landon W12 0NN United Kingdom State (that is, country) of residence: State (that is, country) of nationality. GB GB the States indicated in the Supplemental Box the United States pll designated States except This person is applicant \mathbf{x} all designated States the United States of America of America only for the purposes of:. Name and address: (Family name followed by given name: for a legal entity, full official designation. This person is The address must include postal code and name of country. The country of the address indicated in this Box is the applicant's State (that is, country) of residence if no State of residence is indicated below.) applicant only PUFONG, Boris Tumi X applicant and inventor Gene Expression Technologics Ltd The London BioScience Innovation Centre inventor only (If this check-box is marked. Royal College Street do not fill in below) London Applicant's registration No. with the Office NWI OTU United Kingdom State (that is, country) of residence: State (that is, country) of nationality: GB GB the States indicated in the United States all designated States except This person is applicant all designated States the Supplemental Box the United States of America of America only for the purposes of: Name and address: (Family name followed by given name; for a legal entity, full official designation. This person is The address must include postal code and name of country. The country of the address indicated in this Box Is the applicant's State (that is, country) of residence if no State of residence is indicated below.) applicant only PORTER, Andrew Christopher George X applicant and inventor Gene Targeting Group MRC Clinical Sciences Centre inventor only (If this check-box is marked, Hammersmith Hospital do not fill in below) Du Cane Road Applicant's registration No. with the Office London W12 0NN United Kingdom State (that is, country) of residence: State (that is, country) of nationality: GB GB the United States the States indicated in all designated States except This person is applicant all designated States the Supplemental Box the United States of America of America only for the purposes of. Name and address: (Family name followed by given name: for a legal entity, full official designation. This person is The address must include postal code and name of country. The country of the address indicated in this Bux is the applicant's State (that is, country) of residence if no State of residence is indicated below.) applicant only BULUWELA, Laki Division of Medicine, Cancer Cell Biology Section X applicant and inventor Imperial College School of Medicine, Hammersmith Campus inventor only (If this check-box is marked, Room 601b, Cyclotron Building do not fill in below) Du Cane Road Applicant's registration No. with the Office London W12 0NN United Kingdom State (that is, country) of residence: State (that is, country) of nationality: GB GB the States indicated in the United States all designated States except This person is applicant all designated States the United States of America of America only the Supplemental Box for the purposes of: Further applicants and/or (further) inventors are indicated on another continuation sheet.

Sheet Sheet	t No3	
Continuation of Box No. 111 FURTHER APPLICANT(S) AND/OR	(FURTHER) INVENTOR(S)	
If none of the following sub-baxes is used, this sheet should not be included in		
Name and address: (Family name followed by given name; for a legal entity, full office the address minst include pastal ende and name of country. The country of the address that is the applicant's State (that is, country) of residence of our State of residence to individual VAINIKKA. Satu Gene Expression Technologies Ltd The London BioScience Innovation Centre Royal College Street London NW1 0TU United Kingdom	Indicared in this	rked.
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Further applicants and/or (further) inventors are indicated on another continunt	IAII 4.100-1	

					Sheet No4					
Box	No. Y	designation of states			Mark the applicable check-boxes b	elow; d	ar leost (one must be marked.		
The	followin	ng designations are hereby made under	Rule 4.	9(a):						
Reg	gional Pa	itent								
X	AP	ARIPO Parent: GH Ghana, GM Ga SL Sierra Leone, SZ Swaziland, TZ State which is a Contracting State of specify on dotted line)	United the Har	Republicare Proto	of Tanzania, UG Uganda, ZM Zam	bia, Z' oroseci	W Zimb lion or ti	shwe, and any other realment desired.		
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X	EP	European Patent: AT Austria, BE Belgium, BG Bulgaria, CH & Li Switzerland and Liechtenstein, CY Cyprus, CZ. Czech Republic, DE Germany, DK Denmark, EE Estonia, ES Spain, FI Finland, FR France, GB United Kingdom, GR Greece, IE Ireland, IT Italy, LU Luxembourg, MC Monaco. NL Netherlands, PT Portugal, SE Sweden, SK Slovakia, TR Turkey, and any other State which is a Contracting State of the European Patent Convention and of the PCT OAPI Patent: BF Burkina Faso, BJ Benin, CF Central African Republic, CG Congo, CI Côte d'Ivoire, CM Cameroon,								
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Che	:k-boxes	below reserved for designating States	which l	have bec	ome party to the PCT after issuance	of this	sheet:			
X	vc	St Vincent and Grenadines								
H			H			Ħ				
Prec	autional	ry Designation Statement: In ad	dition	to the	designations made above, the	applica	int also	makes under Rule 4.9(b) all		
arba	e donina	ations which would be nemitted	under	the D	CT except any designation(s) in	idicate	d in t	he Supplemental Box as being		

Precautionary Designation Statement: In addition to the designations made above, the applicant also makes under Rule 4.9(b) all other designations which would be permitted under the PCT except any designation(s) indicated in the Supplemental Box as being excluded from the scope of this statement. The applicant declares that those additional designations are subject to confirmation and that any designation which is not confirmed before the expiration of 15 months from the priority date is to be regarded as withdrawn by the applicant at the expiration of that time limit. (Confirmation (including fees) must reach the receiving Office within the 15-month time limit.)

				Sheet No5		
Box	No. VI PRIOR	TTY CLAIM:				
The	priority of the followi	ng earlier applic	ation(s) is hereby claimed:			
					Where earlier application is	s:
0[6	Filing date curlier application (da)	/month/year)	Number of carlier application	national application; country of Meinber of WTO	regional application:* regional Office	international application: receiving Office
item	(1) I1 Octol	oer 2001	0124391.4	GB		
item	(2)					
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item	(4)					
item	(5)					
	Further priority els	ims are indicate	ed in the Supplemental Box			
The	receiving Office is arlier application was	requested to p	orcpare and transmit to the purpose	he International Bureau a es of this international applic	certified copy of the ear	tier application(s) (only if) identified above as:
	all items	item (1)	item (2)	item (3) ite	em (4) item (5)	Other, see Supplemental Box
• WI Prop	nere the earlier applie erty or one Member o	cation is an AR f the World Tro	de Organization for which i	al least one country party t that earlier application was	filed (Rule 1.10(b)(ii)):	the Protection of Industrial
Bux	No. VII INTERI	NATIONAL S	EARCHING AUTHORIT	Υ		
Choi	ce of International S th, indicate the Author	Searching Autl vity chosen; the	nority (ISA) (if two or mor nwo-letter code may be used	re International Searching A H):	Authorities are competent to	carry out the international
	iest to use results of ching Authority):	earlier search	; reference to that search	i (if an earlier search has b	een carried out by or reque	sied from the International
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Box	No. VIII DECLA	RATIONS				
The l	following declaration in the contraction in the contract in th	s are contained licate in the righ	in Boxes Nos. VIII (i) to (v at column the number of eac) (mark the applicable ch type of declaration);		imber of clarations
	Box No. VIII (i)	Declaration a	s to the identity of the inver	ntor	:	
	Box No. VIII (ii)		s to the applicant's entitlem for and be granted a patent	ient, as at the international fi	iling ;	
	Box No. VIII (iii)		s to the applicant's entitlem the priority of the earlier a	ient, as as the international fi pplication	ling :	
	Box No. VIII (iv)	Declaration of States of Ame		purposes of the designation	of the United :	
	Box No. VIII (v)	Declaration a	s to non-prejudicial disclos	ures or exceptions to lack of	novelty :	

Sheet No ..6..

This international austination contains	UAGE OF FILIN	This international application is accompanied by the following	Numbe
This international application contains: (a) the following number of		item(s) (mark the applicable check-boxes below and Indicate in	of item
sheets in paper form;		right column the number of each item):	
request (including declaration sheets)	: 6	1 fce calculation sheet	:
description (excluding	: 75	2 original separate power of attorney	:
sequence listing part)	•	3. original general power of attorney	:
claims	; 7	4. X copy of general power of attorney; reference number, if any:	: 3
abstract	: 2	5. statement explaining lack of signature	:
(Irawings	: 6	6. priority document(s) identified in Box No. VI as item(s):	:
Sub-total number of sheets:	: 96	7. translation of international application into (language):	:
Sequence listing part of description (actual number of sheets if filed in paper form, whether or not also	: 0	8. separate indications concerning deposited microorganism or other biological material	:
filed in computer readable form; see (b) below)	. •	9. sequence listing in computer readable form (indicate also type and number of carriers (diskette, CD-ROM, CD-R or other))	:
Total number of sheets:	: 96	(i) copy submitted for the purposes of international search under Rule 13 ter only (and not as part of the	
b) Sequence listing part of description fil computer readable form	led in	international application)	:
(i) only (under Section 801(a))(i))	(ii) (only when check-box (b)(i) or (b) (ii) is marked in left column) additional copies including, where applicable,	
(ii)		the copy for the purposes of international search under Rule13ter	:
Type and number of carriers (disket CD-ROM, CD-R or other) on which the sequence listing part is contained (additional december of the contained and decembe	he ditional	(iii) together with relevant statement as to the identity of the copy or copies with the sequence listing part mention in the left column	:
copies to be indicated under item 9(ii), right column):	., m	10. Other (specify):	;
Figure of the drawings which	1	Language of filing of the international application: English	
should accompany the abstract: Box No. X SIGNATURE OF APPL	LICANT AGEN	IT OR COMMON REPRESENTATIVE	
Next in each signature, indicate the name of the p	rerson signing and t	he capacity in which the person signs (if such capacity is not obvious from reading the requ	iest).
		P.J.D. Thomas	
		John S Miles	
	Fo:	receiving Office use only	
Date of actual receipt of the purported international application:	d	2. Drawings	
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Date of timely receipt of the required corrections under PCT Article 11(2):		I I	
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